

AMENDMENTS TO THE CLAIMS

1. (Withdrawn) A flavivirus comprising a hinge region mutation that attenuates the flavivirus.
2. (Withdrawn) The flavivirus of claim 1, wherein the mutation decreases the viscerotropism of the flavivirus.
3. (Withdrawn) The flavivirus of claim 1, wherein the flavivirus comprises a yellow fever virus vaccine strain.
4. (Withdrawn) The flavivirus of claim 1, wherein the flavivirus is a viscerotropic flavivirus selected from the group consisting of Dengue virus, West Nile virus, Wesselsbron virus, Kyasanur Forest Disease virus, and Omsk Hemorrhagic fever virus.
5. (Currently Amended) A vaccine composition comprising a chimeric flavivirus comprising an envelope protein from an attenuated Japanese encephalitis virus, wherein said envelope protein has ~~comprises~~ a substitution ~~hinge~~ region ~~mutation~~ that decreases viscerotropism of the chimeric flavivirus, and said substitution is a reversion to a wild-type sequence at present in an envelope amino acid position 279 selected from the group consisting of amino acids corresponding to amino acids 48-61, 127-131, and 196-283 of a yellow fever virus envelope protein, and the chimeric flavivirus is in the form of a vaccine.

6. (Currently Amended) The vaccine composition chimeric flavivirus of claim 5, wherein the chimeric flavivirus comprises the capsid and non-structural proteins of a first flavivirus virus and the pre-membrane and envelope proteins of said attenuated Japanese encephalitis virus a second flavivirus comprising said substitution that decreases viscerotropism of the chimeric flavivirus and is present in envelope amino acid 279 an envelope protein mutation that attenuates the chimeric flavivirus.

7. (Canceled).

8. (Withdrawn) The flavivirus of claim 6, wherein the second flavivirus is a Dengue virus.

9. (Withdrawn) The flavivirus of claim 8, wherein the Dengue virus is Dengue-1, Dengue-2, Dengue-3, or Dengue-4 virus.

10. (Withdrawn) The flavivirus of claim 1, wherein the mutation is in the hydrophobic pocket of the hinge region of the envelope protein.

11. (Withdrawn) The flavivirus of claim 10, wherein the second flavivirus is a Dengue virus and the mutation is in the lysine at Dengue envelope amino acid position 202 or 204.

12. (Withdrawn) The flavivirus of claim 11, wherein the mutation is a substitution of the lysine.

13. (Withdrawn) The flavivirus of claim 12, wherein the lysine is substituted with arginine.

14. (Withdrawn) A vaccine composition comprising the flavivirus of claim 1 and a pharmaceutically acceptable carrier or diluent.

15. (Withdrawn) A method of inducing an immune response to a flavivirus in a patient, the method comprising administering to the patient the vaccine composition of claim 14.

16. (Withdrawn) The method of claim 15, wherein the patient does not have, but is at risk of developing, infection by the flavivirus.

17. (Withdrawn) The method of claim 15, wherein the patient is infected by the flavivirus.

18. (Withdrawn) A method of producing a vaccine comprising a flavivirus, the method comprising introducing into the flavivirus a mutation that results in decreased viscerotropism.

19. (Withdrawn) The method of claim 18, wherein the mutation is in the hinge region of

the envelope protein of the flavivirus.

20. (Withdrawn) The method of claim 19, wherein the mutation is in the hydrophobic pocket of the envelope protein of the flavivirus.

21. (Withdrawn) A method of identifying a flavivirus vaccine candidate, the method comprising the steps of:

introducing a mutation into the hinge region of the flavivirus; and  
determining whether the flavivirus comprising the hinge region mutation has decreased viscerotropism, as compared with a flavivirus virus lacking the mutation.

22. (Withdrawn) The method of claim 21, wherein the mutation is in the hinge region of the envelope protein of the flavivirus.

23. (Withdrawn) The method of claim 21, wherein the flavivirus is a yellow fever virus.

24. (Withdrawn) The method of claim 21, wherein the flavivirus is a chimeric flavivirus.

25. (Canceled).

26. (Currently Amended) The vaccine composition ~~ehimeric~~ flavivirus of claim 5 [[25]], wherein the substitution hinge region mutation at envelope amino acid position 279 is a

substitution of methionine with lysine.

27. (Currently Amended) The vaccine composition ~~chimeric flavivirus~~ of claim 5 [[25]], wherein the attenuated Japanese encephalitis virus is the JE-SA-14-14-2 strain.

28. (Currently Amended) The vaccine composition ~~chimeric flavivirus~~ of claim 6, wherein the first flavivirus is a yellow fever virus.

29. (Currently Amended) The vaccine composition ~~chimeric flavivirus~~ of claim 28, wherein the yellow fever virus is YF-17D.